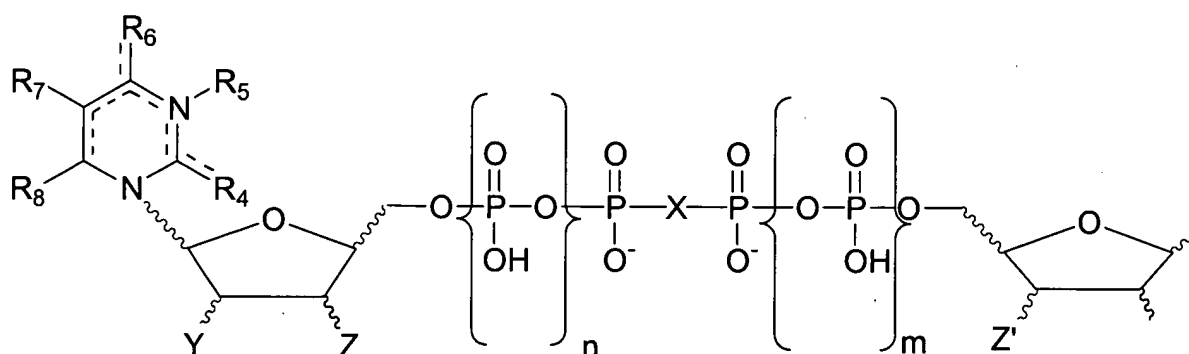


**In the Claims**

1. (Previously Presented) A compound of Formula IIIA:

**Formula IIIA**

wherein:



X is oxygen, methylene, difluoromethylene, imido;

n = 0, 1, or 2;

m = 0, 1, or 2;

n + m = 0, 1, 2, 3, or 4;

B is a purine or a pyrimidine residue linked through the 9- or 1-position, respectively;

Z = OH or N<sub>3</sub>;

Z' = OH or N<sub>3</sub>;

Y = H or OH;

Y' = H or OH;

provided that when Z is N<sub>3</sub>, Y is H or when Z' is N<sub>3</sub>, Y' is H;

R<sub>4</sub> is hydroxy, amino, cyano, aralkoxy, C<sub>1-6</sub> alkoxy, C<sub>1-6</sub> alkylamino, or dialkylamino;

R<sub>5</sub> is hydrogen, acyl, C<sub>1-6</sub> alkyl, phenyloxy, C<sub>1-5</sub> alkanoyl or

absent;

R<sub>6</sub> is oxo, hydroxy, mercapto, C<sub>1-4</sub>alkoxy, C<sub>7-12</sub>arylalkoxy, C<sub>1-6</sub>alkylthio, amino, C<sub>1-5</sub> disubstituted amino, triazolyl, C<sub>1-6</sub>alkylamino or di-C<sub>1-4</sub>alkylamino, where the alkyl groups is optionally linked to form a heterocycle or link to N<sup>3</sup> to form a substituted ring; or

R<sub>5</sub> and R<sub>6</sub> taken together form a 5-membered fused imidazole ring between positions 3 and 4 of the pyrimidine ring, which is optionally substituted on the 4- or 5- positions of the etheno moiety with C<sub>1-4</sub>alkyl, phenyl, or phenyloxy, which themselves are optionally substituted;

R<sub>7</sub> is hydrogen, hydroxy, cyano, nitro, substituted and unsubstituted C<sub>2-8</sub>alkenyl, phenyl, substituted and unsubstituted C<sub>2-8</sub>alkynyl, halogen, CF<sub>3</sub>, substituted and unsubstituted C<sub>1-6</sub>alkyl, allylamino, bromovinyl, ethyl propenoate, propenoic acid; or

R<sub>6</sub> and R<sub>7</sub> taken together form a 5 or 6-membered saturated or unsaturated ring bonded through N or O at R<sub>6</sub>, such ring optionally contain substituents that themselves contain functionalities;

R<sub>8</sub> is hydrogen, amino or di-C<sub>1-4</sub>alkylamino, C<sub>1-4</sub>alkoxy, C<sub>7-12</sub>arylalkoxy, C<sub>1-4</sub>alkylthio, C<sub>7-12</sub>arylalkylthio, carboxamidomethyl, carboxymethyl, methoxy, methylthio, phenoxy or phenylthio; provided that when R<sub>8</sub> is amino or substituted amino, R<sub>7</sub> is hydrogen;

provided that when B = adenine, adenine 1-oxide, or 1,N<sup>6</sup>-ethenoadenine, then:

- (a) R<sub>6</sub> ≠ oxo when R<sub>4</sub> = oxo, Y = Z = OH and R<sub>5</sub> = R<sub>7</sub> = R<sub>8</sub> = H;
- (b) R<sub>7</sub> ≠ Br when R<sub>4</sub> = R<sub>6</sub> = oxo, Y = Z = OH, and R<sub>5</sub> = R<sub>8</sub> = H;

provided that when B = adenine, then:

- (a) R<sub>6</sub> ≠ amino when R<sub>4</sub> = oxo, Y = Z = OH, R<sub>5</sub> is absent, R<sub>7</sub> = R<sub>8</sub> = H, and  
n + m = 0, 1, or 2;
- (b) R<sub>7</sub> ≠ CH<sub>3</sub> when R<sub>4</sub> = R<sub>6</sub> = oxo, Y = H, Z = OH, and R<sub>5</sub> = R<sub>8</sub> = H;
- (c) R<sub>7</sub> ≠ F when R<sub>4</sub> = R<sub>6</sub> = oxo, Y = H, Z = OH, R<sub>5</sub> = R<sub>8</sub> = H and n + m = 2;

provided that when B = thymine, Y' = H and Z' = N<sub>3</sub>; then R<sub>7</sub> ≠ F, when R<sub>4</sub> = R<sub>6</sub> = oxo, Y = OH, Z = OH, R<sub>5</sub> = R<sub>8</sub> = H, and n + m = 0;

provided that when B = thymine, Y' = H and Z' = N<sub>3</sub>; then R<sub>7</sub> ≠ CH<sub>3</sub> when R<sub>4</sub> = R<sub>6</sub> = oxo, Y = H, Z = N<sub>3</sub>, R<sub>5</sub> = R<sub>8</sub> = H, and n + m = 0;

provided that when B = guanine, then:

- (a)  $R_6 \neq \text{oxo}$  when  $R_4 = \text{oxo}$ ,  $Y = Z = \text{OH}$ ,  $R_5 = R_7 = R_8 = \text{H}$  and  $n + m = 1$  or  $2$ ;
- (b)  $R_6 \neq \text{amino}$  when  $R_4 = \text{oxo}$ ,  $Y = Z = \text{OH}$ ,  $R_5$  is absent,  $R_7 = R_8 = \text{H}$ ,  $n+m=1$  or  $2$ ;

provided that when B is uridine, or 5-Br-uridine, then

- (a)  $R_6 \neq \text{oxo}$  when  $R_4 = \text{oxo}$ ,  $Y = Z = \text{OH}$  and  $R_6 = R_7 = R_8 = \text{H}$ ;
- (b)  $R_7 \neq \text{Br}$  when  $R_4 = R_6 = \text{oxo}$ ,  $Y = Z = \text{OH}$ , and  $R_5 = R_8 = \text{H}$ ;

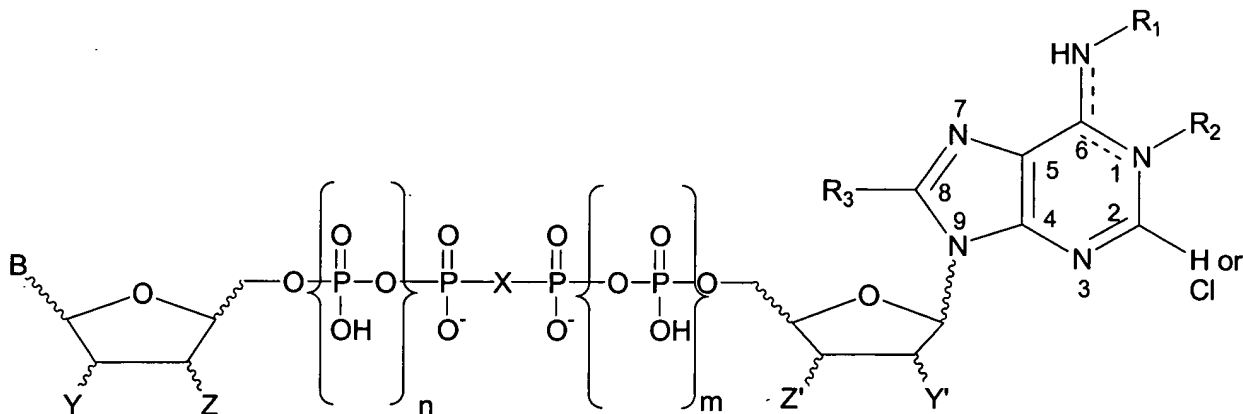
provided that when B is 5-FU, then  $R_7 \neq \text{F}$ , when  $R_4 = R_6 = \text{oxo}$ ,  $Y = \text{H}$ ,  $Z = \text{OH}$ ,  $R_5 = R_8 = \text{H}$ , and  $n + m = 0$ ;

provided that when B is cytosine, then  $R_6 \neq \text{amino}$ , when  $R_4 = \text{oxo}$ ,  $Y = Z = \text{OH}$ ,  $R_5$  is absent,  $R_7 = R_8 = \text{H}$ , and  $n + m = 1$ , or  $2$ ; and

provided that when B is cytosine, then  $R_6 \neq \text{oxo}$ , when  $R_4 = \text{oxo}$ ,  $Y = Z = \text{OH}$  and  $R_6 = R_7 = R_8 = \text{H}$ , and  $n + m = 2$ .

2. (Original) A compound according to Formula IIA:

FORMULA IIA



wherein:

X is oxygen, methylene, difluoromethylene, imido;

n = 0, 1, or 2;

m = 0, 1, or 2;

n + m = 0, 1, 2, 3, or 4;

B is a purine residue linked through the 9- position;

Z = OH or N<sub>3</sub>;

Z' = OH or N<sub>3</sub>;

Y = H or OH;

Y' = H or OH;

provided that when Z is N<sub>3</sub>, Y is H or when Z' is N<sub>3</sub>, Y' is H;

R<sub>1</sub> is H, C<sub>1-8</sub>alkyl, phenyl or phenyloxy, optionally substituted with halogen, hydroxy, C<sub>1-4</sub>alkoxy, C<sub>1-4</sub>alkyl, C<sub>6-10</sub>aryl, carboxy, cyano, nitro, sulfonamido, sulfonate, phosphate, sulfonic acid, amino or substituted amino, wherein the amino is singly or doubly substituted by a C<sub>1-4</sub> alkyl and when doubly substituted, the alkyl groups are optionally linked to form a heterocycle; or A(C<sub>1-6</sub>alkyl)CONH(C<sub>1-6</sub>alkyl)B wherein A and B are amino, mercapto, hydroxy or carboxyl;

R<sub>2</sub> is O or is absent; or

R<sub>1</sub> and R<sub>2</sub> taken together forms a 5-membered fused imidazole ring, which is optionally substituted on the 4- or 5- positions of the etheno moiety with C<sub>1-4</sub>alkyl, phenyl or phenyloxy, optionally substituted with halogen, hydroxy, C<sub>1-4</sub>alkoxy, C<sub>1-4</sub>alkyl, C<sub>6-10</sub>aryl, arylalkyl, carboxy, cyano, nitro, sulfonamido, sulfonate, phosphate, sulfonic acid, amino or substituted amino, wherein the amino is singly or doubly substituted by a C<sub>1-4</sub> alkyl and when doubly substituted, the alkyl groups is optionally linked to form a heterocycle; and

R<sub>3</sub> is H, C<sub>1-8</sub>alkyl, phenyl or phenyloxy, optionally substituted with halogen, hydroxy, C<sub>1-4</sub>alkoxy, C<sub>1-4</sub>alkyl, C<sub>6-10</sub>aryl, carboxy, cyano, nitro, sulfonamido, sulfonate,

phosphate, sulfonic acid, amino or substituted amino, wherein the amino is singly or doubly substituted by a C<sub>1-4</sub> alkyl and when doubly substituted, the alkyl groups is optionally linked to form a heterocycle; C<sub>7-12</sub>arylalkyl; C<sub>1-4</sub>alkylamino, phenylamino, C<sub>7-12</sub>arylalkylamino, C<sub>1-4</sub>alkoxy, or C<sub>7-12</sub>arylalkyloxy; C<sub>1-4</sub>alkylthio, phenylthio, C<sub>7-12</sub>arylalkylthio, or -A(C<sub>1-6</sub>alkyl)CONH(C<sub>1-6</sub>alkyl)B- wherein A and B are independently amino, mercapto, hydroxy or carboxyl;

provided that R<sub>1</sub> ≠ H, when X is oxygen, methylene, or difluoromethylene, Y is OH, B is adenine, R<sub>2</sub> is absent, and R<sub>3</sub> is hydrogen;

provided that R<sub>1</sub> ≠ H, when n + m = 2, X is oxygen, Y is OH, B is adenine, R<sub>2</sub> is absent, and R<sub>3</sub> is bromo, or 6-aminohexyl;

provided that R<sub>1</sub> ≠ H, when n + m = 2, X is oxygen, Y is H, B is adenine, R<sub>2</sub> is absent, and R<sub>3</sub> is H;

provided that R<sub>2</sub> ≠ O, when n + m = 2, X is oxygen, Y is OH, R<sub>1</sub> = R<sub>3</sub> = H, and B is adenine, adenine 1-oxide, or 1,N<sup>6</sup>-ethenoadenine;

provided that R<sub>1</sub> and R<sub>2</sub> do not form a 5-membered fused imidazole ring, when n + m = 2, X is oxygen, Y is OH, R<sub>3</sub> is H, and B is adenine, adenine 1-oxide, or ethenoadenine.

3. (Original) The compound according to Claim 1 or 2, wherein the ribosyl moieties are in the D- configuration.

4. (Original) The compound according to Claim 1 or 2, wherein the ribosyl moieties are in the L- configuration.

5. (Previously Presented) A pharmaceutical composition comprising a compound of Formula IIIA or IIA as described in Claim 1 or 2, or a pharmaceutically acceptable salt thereof together with a pharmaceutically acceptable carrier therefor.

6. (Previously Presented) A method of treating chronic obstructive pulmonary diseases in a mammal by administering an effective chronic obstructive pulmonary disease treatment amount of a compound of Formula IIIA or IIA as described in Claim 1 or 2.

7. (Previously Presented) A method of treating sinusitis, otitis media or nasolacrimal duct obstruction in a mammal by administering an effective mucus secretion clearing amount of a compound of Formula IIIA or IIA as described in Claim 1 or 2.

8. (Previously Presented) A method of treating dry eye in a mammal by administering an effective dry eye treatment amount of a compound of Formula III A or IIA as described in Claim 1 or 2.

9. (Previously Presented) A method of treating retinal detachment in a mammal by administering an effective retinal detachment treatment amount of a compound of Formula IIIA or IIA as described in Claim 1 or 2.

10. (Currently Amended) A method of facilitating sputum induction in a mammal by administering an effective amount of a compound of Formula ~~IA or IB~~ IIIA or IIA as described in Claim 1 or 2, ~~effective~~ to facilitate sputum induction.

11. (Currently Amended) A method of facilitating expectoration in a mammal by administering an effective amount of a compound of Formula ~~IA or IB~~ IIIA or IIA as described in Claim 1 or 2, ~~effective~~ to facilitating expectoration.

12. (New) A method of treating cystic fibrosis in a mammal by administering an effective amount of a compound of Formula IIIA or IIA as described in Claim 1 or 2 to treat cystic fibrosis.

13. (New) The method according to Claim 12, wherein said compound is P<sup>1</sup>-(2'-deoxycytidine 5')-P<sup>4</sup>-( uridine 5')tetraphosphate.